SEQUENCE LISTING

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<110> Crooke, Stanley T Lima, Walter Wu, Hongjiang

<120> Methods of Using Mammalian RNase H and Compositions Thereof

<130> ISPH-0520

<140> US 09/781,712

<141> 2001-02-12

<150> US 60/067,458

<151> 1997-12-04

<150> US 09/203,716

<151> 1998-12-02

<150> US 09/343,809

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<151> 2000-10-06

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Ile Cys Tyr Cys Pro Leu Pro Arg Leu Ala Asp Leu Glu Ala Leu Leu 50 55 60

Val Ala Asp Ser Leu Thr Leu Leu Glu Ser Glu Arg Glu Arg Leu Phe 65 70 75 80

Ala Leu Met Glu Asp Thr Asp Phe Val Gly Trp Ala Leu Asp Val Leu 85 90 95

Ser Pro Asn Leu Ile Ser Thr Ser Met Leu Gly Trp Val Leu Tyr Asn

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Leu Asp Gln Gly Val Asn Val Thr Gln Val Phe Val Asp Thr Val Gly 130 135 140

Glu Val Thr Val Leu Ala Leu Ala Asp Ala Leu Tyr Pro Val Val Ser 165 170 175

Ala Ala Ser Ile Cys Ala Leu Val Ala Arg Asp Gln Ala Val Leu Leu 180 185 190

Trp Gln Phe Val Glu Leu Leu Gln Asp Leu Asp Thr Asp Tyr Gly Ser 195 200 205

Gly Tyr Pro Asn Asp Pro Leu Thr Leu Ala Trp Leu Leu Glu His Val 210 215 220

Glu Pro Val Phe Gly Phe Pro Gln Phe Val Arg Phe Ser Trp Arg Thr 225 230 235 240

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p Glu Asp 245 250 255

Ser Ala Ser Glu Asn Gln Glu Gly Leu Arg Leu Ile Thr Ser Tyr Phe 260 265 270

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Ile Cys Tyr Cys Pro Leu Ser Arg Leu Ala Asp Leu Glu Ala Leu Leu 50 55 60

Val Ala Asp Ser Leu Thr Leu Thr Glu Asn Glu Arg Glu Arg Leu Phe 65 70 75 80

Ala Leu Met Glu Glu Asp Gly Asp Phe Val Gly Trp Ala Leu Asp Val 85 90 95

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Asn Leu Gly Val Asp Asp Ser Leu Ala Leu Asn Glu Ala Leu Arg Glu 65 70 75 80

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Ala Tyr Ala Leu Arg Cys Leu Ser Pro Glu Leu Ile Ser Cys Ser Met

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Leu Leu Arg Gln Leu Tyr Ser Leu Asn Glu Val Ser His Glu Ala Ala 120

Ile Thr Leu Ile Arg Asp Ala Leu Ala Cys Asn Val Asn Val Val Glu 135 140

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Ser Leu Phe Pro Ile Val Ser Ala Ala Ser Ile Ala Ala Leu Val Thr

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Leu Val Arg Ser Ser Trp Leu Thr Ala Ser Thr Ile Val Glu Leu Arg 245

Cys Val Pro Gly Ser Trp Glu Asp Asp Glu Glu Glu Gly Leu Ser Gln 260 265

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Pro Met Val Tyr Ala Val Ala Tyr Ser Thr Gln Leu Tyr Gln Asp Glu 50 55 60

Thr Ile Ile Pro Asn Tyr Glu Phe Asp Asp Ser Leu Leu Leu Thr Asp 65 70 75 80

Pro Ile Arg Arg Met Leu Phe Ser Leu Ile Tyr Gln Asp Asn Glu Glu 85 90 95

Leu Thr Gln Ile Gly Tyr Ala Thr Thr Cys Ile Thr Pro Leu Asp Ile 100 105 110

Ser Arg Gly Met Ser Leu Phe Pro Pro Thr Arg Asn Tyr Asn Leu Asn 115 120 125

Glu Gln Ala His Asp Val Thr Met Ala Leu Ile Asp Gly Val Ile Leu 130 135 140

Gln Asn Val Leu Leu Ser His Val Tyr Val Asp Thr Val Gly Pro Pro 145 150 155 160

Ala Ser Tyr Gln Leu Leu Glu Gln Arg Phe Pro Gly Val Leu Phe 165 170 175

Thr Val Ala Leu Leu Ala Asp Ser Leu Tyr Cys Met Val Ser Val Ala 180 185 190

Ser Val Val Ala Leu Val Thr Arg Asp Ile Leu Val Glu Ser Leu Leu 195 200 205

Arg Asp Pro Asp Glu Ile Leu Gly Ser Gly Tyr Pro Ser Asp Pro Leu 210 215 220

Thr Val Ala Trp Leu Leu Arg Asn Gln Thr Ser Leu Met Gly Trp Pro 225 230 235 240

Ala Asn Met Val Arg Phe Ser Trp Gln Thr Cys Gln Thr Leu Leu Asp 245 250 255

Asp Ala Ser Leu Asn Ser Ile Pro Ile Leu Trp Glu Glu Gln Tyr Met 260 265 270

Asp Ser Arg Leu Asn Ala Ala Gln Leu Thr Leu Gln Leu Gln Leu Gln 275 280 285

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Leu Ser Glu Leu Arg Arg Leu Ala Leu Tyr Glu Glu Ile Leu Glu Leu 50 55 60

Ala Leu Ser Trp Ser Leu Gly Arg Ala Glu Pro His Glu Ile Asp Glu 65 70 75 80

Leu Asn Ile Leu His Ala Thr Met Leu Ala Met Gln Arg Ala Val Ala 85 90 95

Gly Leu His Ile Ala Pro Glu Tyr Val Leu Ile Asp Gly Asn Arg Cys 100 105 110

Pro Leu Leu Pro Met Pro Ala Met Ala Val Val Leu Gly Asp Ser Arg 115 120 125

Val Pro Glu Ile Ser Ala Ala Ser Ile Leu Ala Leu Val Thr Arg Asp 130 135 140 Gln His Leu Gly Tyr Pro Thr Ala Phe His Leu Glu Leu Leu Ala Glu 165 170 175

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Ala Leu Gly Leu Ala Ser 195

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Gln Val Asp Arg Phe Pro Ala Ala Arg Phe Leu Leu Phe Ala Thr Glu 50 55 60

Asp Glu Ala Trp Ala Phe Val Arg Leu Ser Ala Ser Pro Glu Val Ser 65 70 75 80

Glu Gly His Glu Asn Gln His Gly Gln Glu Ser Glu Ala Leu Pro Gly 85 90 95

Leu Arg Leu Arg Glu Pro Leu Asp Gly Asp Gly His Glu Ser Ala Gln
100 105 110

Pro Tyr Ala Leu His Met Leu Pro Ser Val Glu Pro Ala Pro Pro Val 115 120 125

Ser Arg Asp Thr Phe Ser Tyr Met Gly Asp Phe Val Val Tyr Thr 130 135 140

Gly Val Tyr Trp Gly Pro Gly His Pro Leu Asn Val Gly Ile Arg Leu 165 170 175

Pro Gly Arg Gln Thr Asn Gln Arg Ala Glu Ile His Ala Ala Cys Leu 180 185 190

Ala Ile Glu Gln Ala Leu Thr Gln Asn Ile Asn Leu Leu Val Leu Tyr 195 200 205

Thr Asp Ser Met Phe Thr Ile Asn Gly Ile Thr Asn Trp Val Gln Gly 210 215 220

Trp Leu Leu Asn Gly Trp Leu Thr Ser Ala Gly Leu Glu Val Ile Asn 225 230 235 240

Leu Glu Asp Phe Val Ala Leu Glu Arg Leu Thr Gln Gly Met Asp Ile 245 250 255

Gln Trp Met His Val Pro Gly His Ser Gly Phe Ile Gly Asn Glu Glu 260 265 270

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Gln Val Asp Arg Phe Pro Ala Ala Arg Phe Leu Leu Phe Ala Thr Glu 50 60

Asp Glu Ala Trp Ala Phe Val Arg Leu Ser Ala Ser Pro Glu Val Ser 65 70 75 80

Glu Gly His Glu Asn Gln His Gly Gln Glu Ser Glu Ala Leu Ala Ser

Leu Arg Leu Arg Glu Pro Leu Asp Gly Asp Gly His Glu Ser Ala Glu 100 105

Pro Tyr Ala Leu His Met Leu Pro Ser Val Glu Pro Ala Pro Pro Val

Ser Arg Asp Thr Phe Ser Tyr Met Gly Asp Phe Val Val Val Tyr Thr 135 130

Asp Gly Cys Cys Ser Ser Asn Gly Arg Arg Pro Arg Ala Gly Ile 145 150 155

Gly Val Tyr Trp Gly Pro Gly His Pro Leu Asn Val Gly Ile Arg Leu 165 170

Pro Gly Arg Gln Thr Asn Gln Arg Ala Glu Ile His Ala Ala Cys Leu 180 185 190

Ala Ile Glu Gln Ala Leu Thr Gln Asn Ile Asn Leu Leu Val Leu Tyr

Thr Asp Ser Met Phe Thr Ile Asn Gly Ile Thr Asn Trp Val Gln Gly 215

Trp Leu Leu Asn Gly Trp Leu Thr Ser Ala Gly Leu Glu Val Ile Asn 225 230 235

Leu Glu Asp Phe Val Ala Leu Glu Arg Leu Thr Gln Gly Met Asp Ile 245

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Arg Gly Arg Leu Thr Gly Val Phe Leu Thr Trp Asn Glu Cys Arg Ala 35 40 45

Gln Val Asp Arg Phe Pro Ala Ala Arg Phe Leu Leu Phe Ala Thr Glu 50 55 60

Asp Glu Ala Trp Ala Phe Val Arg Leu Ser Ala Ser Pro Glu Val Ser 65 70 75 80

Glu Gly His Glu Asn Gln His Gly Gln Glu Ser Glu Ala Leu Ala Ser 85 90 95

Leu Arg Leu Arg Glu Pro Leu Asp Gly Asp Gly His Glu Ser Ala Glu 100 105 110

Pro Tyr Ala Leu His Met Leu Pro Ser Val Glu Pro Ala Pro Pro Val 115 120 125

Ser Arg Asp Thr Phe Ser Tyr Met Gly Asp Phe Val Val Tyr Thr 130 135 140

Asp Gly Cys Cys Ser Ser Asn Gly Arg Arg Pro Arg Ala Gly Ile 145 150 155 160

Gly Val Tyr Trp Gly Pro Gly His Pro Leu Asn Val Gly Ile Arg Leu 165 170 175

Pro Gly Arg Gln Thr Asn Gln Arg Ala Glu Ile His Ala Ala Cys Leu 180 185 190

Ala Ile Glu Gln Ala Leu Thr Gln Asn Ile Asn Leu Leu Val Leu Tyr 195 200 205

Thr Asp Ser Met Phe Thr Ile Asn Gly Ile Thr Asn Trp Val Gln Gly 210 215 220

Trp Leu Leu Asn Gly Trp Leu Thr Ser Ala Gly Leu Glu Val Ile Asn 225 230 235 240

Leu Glu Asp Phe Val Ala Leu Glu Arg Leu Thr Gln Gly Met Asp Ile

245 250 255

Gln Trp Met His Val Pro Gly His Ser Gly Phe Ile Gly Asn Glu Glu 260 265 270

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Arg Gly Arg Leu Thr Gly Val Phe Leu Thr Trp Asn Glu Cys Arg Ala 35 40 45

Gln Val Asp Arg Phe Pro Ala Ala Arg Phe Leu Leu Phe Ala Thr Glu 50 55 60

Asp Glu Ala Trp Ala Phe Val Arg Leu Ser Ala Ser Pro Glu Val Ser 65 70 75 80

Glu Gly His Glu Asn Gln His Gly Arg Glu Ser Glu Ala Leu Ala Ser 85 90 95

Leu Arg Leu Arg Glu Pro Leu Asp Gly Asp Gly His Glu Ser Ala Glu 100 105 110

Pro Tyr Ala Leu His Met Leu Pro Ser Val Glu Pro Ala Pro Pro Val 115 120 125

Ser Arg Asp Thr Phe Ser Tyr Met Gly Asp Phe Val Val Val Tyr Thr 130 135 140

Gly Val Tyr Trp Gly Pro Gly His Pro Leu Asn Val Gly Ile Arg Leu 165 170 175 Pro Gly Arg Gln Thr Asn Gln Arg Ala Glu Ile His Ala Ala Cys Leu 180 185 190

Ala Ile Glu Gln Ala Leu Thr Gln Asn Ile Asn Leu Leu Val Leu Tyr 195 200 205

Thr Asp Ser Met Phe Thr Ile Asn Gly Ile Thr Asn Trp Val Arg Gly 210 215 220

Trp Leu Leu Asn Gly Trp Leu Thr Ser Ala Gly Leu Glu Val Ile Asn 225 230 235 240

Leu Glu Asp Phe Val Ala Leu Glu Arg Leu Thr Gln Gly Met Asp Ile 245 250 255

Gln Trp Met His Val Pro Gly His Ser Gly Phe Ile Gly Asn Glu Glu 260 265 270

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20 25 30

Val Asp Glu Ala Gly Arg Gly Pro Val Leu Gly Pro Met Val Tyr Ala 35 40 45

Ile Cys Tyr Cys Pro Leu Pro Arg Leu Ala Asp Leu Glu Ala Leu Leu 50 55 60

Val Ala Asp Ser Leu Thr Leu Leu Glu Ser Glu Arg Glu Arg Leu Phe 70 75 80

Ala Leu Met Glu Asp Thr Asp Phe Val Gly Trp Ala Leu Asp Val Leu 85 90 95

Ser Pro Asn Leu Ile Ser Thr Ser Met Leu Gly Arg Val Leu Tyr Asn

100 105 110

Leu Asn Ser Leu Ser His Asp Thr Ala Thr Gly Leu Ile Gln Tyr Ala 115 120 125

Leu Asp Gln Gly Val Asn Val Thr Gln Val Phe Val Asp Thr Val Gly 130 135 140

Glu Val Thr Val Leu Ala Leu Ala Asp Ala Leu Tyr Pro Val Val Ser 165 170 175

Ala Ala Ser Ile Cys Ala Leu Val Ala Arg Asp Gln Ala Val Leu Leu 180 185 190

Trp Gln Phe Val Glu Leu Leu Gln Asp Leu Asp Thr Asp Tyr Gly Ser 195 200 205

Gly Tyr Pro Asn Asp Pro Leu Thr Leu Ala Trp Leu Leu Glu His Val 210 $\,$ 215 $\,$ 220 $\,$

Glu Pro Val Phe Gly Phe Pro Gln Phe Val Arg Phe Ser Trp Arg Thr 225 230 235 240

Ala Gln Thr Ile Leu Glu Leu Glu Ala Glu Asp Val Ile Trp Glu Asp 245 250 255

Ser Ala Ser Glu Asn Gln Glu Gly Leu Arg Leu Ile Thr Ser Tyr Phe 260 265 270

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Leu Glu Arg Gly Leu Glu Ser Ala Thr Ser Leu 290 295

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- Arg Gly Arg Arg Thr Gly Val Phe Leu Ser Trp Ser Glu Cys Leu Ala 35 40 45
- Gln Val Asp Arg Phe Pro Ala Ala Arg Phe Leu Leu Phe Ala Thr Glu 50 55 60
- Asp Glu Ala Trp Ala Phe Val Arg Ser Ser Ser Ser Pro Asp Gly Ser 65 70 75 80
- Leu Gly Gln Glu Ser Ala His Glu Gln Leu Ser Gln Ala Leu Thr Ser 85 90 95
- Leu Arg Pro Arg Glu Pro Leu Gly Glu Glu Glu Leu Pro Glu Pro 100 105 110
- Gly Pro Leu His Thr Arg Gln Asp Thr Glu Pro Ala Ala Val Val Ser 115 120 125
- Leu Asp Thr Phe Ser Tyr Met Gly Glu Ser Val Ile Val Tyr Thr Asp 130 135 140
- Val Tyr Trp Gly Pro Gly His Pro Leu Asn Val Gly Ile Arg Leu Pro 165 170 175
- Gly Arg Gln Thr Asn Gln Arg Ala Glu Ile His Ala Ala Cys Leu Ala 180 185 190
- Ile Met Gln Ala Leu Ala Gln Asn Ile Ser Leu Leu Val Leu Tyr Thr 195 200 205
- Asp Ser Met Phe Thr Ile Asn Gly Ile Thr Asn Trp Val Gln Gly Trp 210 215 220
- Leu Leu Asn Gly Trp Arg Thr Ser Thr Gly Leu Asp Val Ile Asn Leu 225 230 235 240
- Glu Asp Phe Met Glu Leu Asp Glu Leu Thr Gln Gly Met Asp Ile Gln 245 250 255

Trp Met His Ile Pro Gly His Ser Gly Phe Val Gly Asn Glu Glu Ala 260 265 270

Asp Arg Leu Ala Arg Glu Gly Ala Leu Gln Ser Glu Asp 275 280 285

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